

✓ fifth line from last, after "triglycerides"
insert a comma -- , --.

N/E Page 16, seventh line after Table 10, correct the
spelling of -- disease.--.

✓
Page 17, second line, after "to" insert -- the --;
eighth line from last, after "to" insert
-- the --; and
third line from last, after "to" insert -- the
--.

✓
Page 18, ninth line from last, change "carrier" to
--carriers--.

✓
Page 19, first line after the Example table, change
"weighted" to read -- weighed --; and
next to last line, delete the comma.

IN THE CLAIMS:

Cancel Claims 1-15.

Please enter the following new claims:

B¹ --'16. A method for elevating the HDL
cholesterol level in the serum of a human patient, which
comprises administering to the patient a pharmaceutical
composition in which the active ingredients consist
essentially of a mixture of fatty acids of which at least 80%
by weight is comprised of a combination of (all-Z omega-3)-

Contd.
B1

5,8,11,14,17-eicosapentaenoic acid (EPA) and (all-Z omega-3)-4,7,10,13,16,19-docosahexaenoic acid (DHA) in a weight ratio of EPA:DHA of from 1:2 to 2:1, said composition being administered in amounts providing a daily dosage of 1 to 10 grams of said mixture of fatty acids.

2₁₇. The method of claim ¹~~16~~, wherein at least 85% by weight of the mixture of fatty acids is comprised of long chain omega-3 fatty acids.

3₁₈. The method of claim ²~~17~~, wherein the EPA constitutes 40 to 60% by weight of the mixture of fatty acids and the DHA constitutes 25 to 45% by weight of the mixture of fatty acids.

4₁₉. The method of claim ³~~18~~, wherein the EPA and DHA are present in the composition in an EPA:DHA weight ratio of from 1:1 to 2:1.

9₂₀. The method of claim ⁴~~19~~, wherein at least 4.5% by weight of the mixture of fatty acids is comprised of fatty acids other than EPA and DHA that have 20, 21, or 22 carbon atoms.

5₂₁. The method of claim ⁴~~19~~, wherein at least 3% by weight of the mixture of fatty acids is comprised of omega-3 fatty acids other than EPA and DHA that have 20, 21, or 22 carbon atoms.

Contd.
B1

⁶
~~22~~. The method of claim ⁴~~19~~, wherein at least 1% by weight of the mixture of fatty acids is comprised of (all-Z omega-3)-6,9,12,15,18-heneicosapentaenoic acid.

^{9 5 6}
¹⁰~~23~~. The method of any of claims ~~20~~, ~~21~~, or ~~22~~, wherein the fatty acids are present in the composition in esterified form.

^{9 5 6}
¹¹~~24~~. The method of any of claims ~~20~~, ~~21~~, or ~~22~~, wherein the fatty acids are present in the composition in ethyl ester form.

^{9 5 6}
¹²~~25~~. The method of any of claims ~~20~~, ~~21~~, or ~~22~~, wherein the fatty acids are present in the composition in salt form.

^{9 5 6}
¹³~~26~~. The method of any of claims ~~20~~, ~~21~~, or ~~22~~, wherein the fatty acids are present in the composition in the free acid form.

⁶
⁷~~27~~. The method of claim ~~22~~, wherein at least 85% by weight of the fatty acid content of the composition is comprised of the combination of EPA and DHA, and the fatty acids are present in the composition in ethyl ester form.

⁴
⁸~~28~~. The method of claim ~~19~~, wherein the composition is administered orally.

[
29. A method for the treatment or prophylaxis of hypertension in an adult human patient, which comprises

Contd.

B1

administering to the patient, on a daily basis, an effective amount of a pharmaceutical composition in which the active ingredients consist essentially of a mixture of fatty acids of which at least 80% by weight is comprised of a combination of (all-Z omega-3)-5,8,11,14,17-eicosapentaenoic acid (EPA) and (all-Z omega-3)-4,7,10,13,16,19-docosahexaenoic acid (DHA) in a weight ratio of EPA:DHA of from 1:2 to 2:1.

30. The method of claim 29, wherein at least 85% by weight of the mixture of fatty acids is comprised of long chain omega-3 fatty acids.

31. The method of claim 30, wherein the EPA constitutes 40 to 60% by weight of the mixture of fatty acids and the DHA constitutes 25 to 45% by weight of the mixture of fatty acids.

32. The method of claim 31, wherein the EPA and DHA are present in the composition in an EPA:DHA weight ratio of from 1:1 to 2:1.

33. The method of claim 32, wherein at least 3% by weight of the mixture of fatty acids is comprised of omega-3 fatty acids other than EPA and DHA that have 20, 21, or 22 carbon atoms.

34. The method of claim 33, wherein at least 90% by weight of the composition is comprised of long chain, polyunsaturated, omega-3 fatty acids.

35. The method of claim 34, wherein at least 1% by weight of the mixture of fatty acids is comprised of (all-Z omega-3)-6,9,12,15,18-heneicosapentaenoic acid.

36. The method of any of claims 33, 34, or 35, wherein the fatty acids are present in the composition in esterified form.

37. The method of any of claims 33, 34, or 35, wherein the fatty acids are present in the composition in ethyl ester form.

38. The method of any of claims 33, 34, or 35, wherein the fatty acids are present in the composition in the free acid form.

39. The method of claim 35, wherein at least 85% by weight of the fatty acid content of the composition is comprised of the combination of EPA and DHA, and the fatty acids are present in the composition in ethyl ester form.

40. The method of claim 32, wherein the composition is administered orally.

41. A method for the treatment or prophylaxis of multiple risk factors for cardiovascular diseases, which comprises administering to a patient a mixed-fatty acids composition in which

Contd.

B1

a) at least 80% by weight of the composition is comprised of omega-3 fatty acids,

b) at least 80% by weight of the total fatty acid content of the composition is comprised of a combination of (all-Z omega-3)-5,8,11,14,17-eicosapentaenoic acid (EPA) and (all-Z omega-3)-4,7,10,13,16,19-docosahexaenoic acid (DHA) in a weight ratio of EPA:DHA of from 1:2 to 2:1, and

c) the fatty acids are in admixture with a pharmaceutically acceptable carrier.

42. A method for the treatment or prophylaxis of multiple risk factors for cardiovascular diseases in an adult human patient, which comprises orally administering to the patient a pharmaceutical composition in which the active ingredients consist essentially of a mixture of fatty acids of which at least 80% by weight is comprised of a combination of (all-Z omega-3)-5,8,11,14,17-eicosapentaenoic acid (EPA) and (all-Z omega-3)-4,7,10,13,16,19-docosahexaenoic acid (DHA) in a weight ratio of EPA:DHA of from 1:2 to 2:1, said composition being administered in amounts providing a daily dosage of 1 to 10 grams of said mixture of fatty acids.

43. The method of claim 42, wherein at least 85% by weight of the mixture of fatty acids is comprised of long chain omega-3 fatty acids.

44. The method of claim 43, wherein the EPA constitutes 40 to 60% by weight of the mixture of fatty acids

Contd.

81 and the DHA constitutes 25 to 45% by weight of the mixture of fatty acids.

45. The method of claim 44, wherein the EPA and DHA are present in the composition in an EPA:DHA weight ratio of from 1:1 to 2:1.

46. The method of claim 45, wherein at least 4.5% by weight of the mixture of fatty acids is comprised of fatty acids other than EPA and DHA that have 20, 21, or 22 carbon atoms.

47. The method of claim 45, wherein at least 3% by weight of the mixture of fatty acids is comprised of omega-3 fatty acids other than EPA and DHA that have 20, 21, or 22 carbon atoms.

48. The method of claim 45, wherein at least 1% by weight of the mixture of fatty acids is comprised of (all-Z omega-3)-6,9,12,15,18-heneicosapentaenoic acid.

49. The method of any of claims 46, 47, or 48, wherein the fatty acids are present in the composition in esterified form.

50. The method of any of claims 46, 47, or 48, wherein the fatty acids are present in the composition in ethyl ester form.

Contd
B1

51. The method of any of claims 46, 47, or 48, wherein the fatty acids are present in the composition in salt form.

52. The method of any of claims 46, 47, or 48, wherein the fatty acids are present in the composition in the free acid form.

14/ 53. A pharmaceutical mixed-fatty-acids composition in which

a) at least 80% by weight of the composition is comprised of a combination of (all-Z omega-3)-5,8,11,14,17-eicosapentaenoic acid (EPA) and (all-Z omega-3)-4,7,10,13,16,19-docosahexaenoic acid (DHA) in a weight ratio of EPA:DHA of from 1:2 to 2:1 and

b) (all-Z omega-3)-6,9,12,15,18-heneicosapentaenoic acid is present in an amount of at least one percent by weight.

15/ 54. The composition of claim 53, wherein at least 85% by weight of the composition is comprised of long chain omega-3 fatty acids.

16/ 55. The composition of claim 54, wherein the EPA constitutes 40 to 60% by weight of the composition and the DHA constitutes 25 to 45% by weight of the composition.

Contd.
81

¹⁷
~~56~~. The composition of claim ¹⁶~~55~~, wherein C 20:4 omega-6 fatty acid constitutes at least one percent by weight of the composition.

¹⁸ ¹⁹
~~57~~. The composition of claim ¹⁷~~56~~, wherein C 22:5 omega-3 fatty acid constitutes at least one percent by weight of the composition.

¹⁹ ²⁰
~~58~~. The composition of claim ¹⁶~~55~~, wherein the (all-Z omega-3)-6,9,12,15,18-heneicosapentaenoic acid is present in an amount of from 1 to 4% by weight.

²¹
~~59~~. The composition of claim ¹⁹ ²⁰~~58~~, wherein C 22:5 omega-3 fatty acid constitutes 1 to 3% by weight of the composition.

¹⁶ ²³
~~55~~. The composition of any of claims ¹⁴ ¹⁵~~53~~, ~~54~~, or ~~55~~, wherein the EPA and DHA are present in an EPA:DHA weight ratio of from 1:1 to 2:1.

¹⁹ ²⁰ ²⁰ ¹⁸
~~58~~. The composition of any of claims ¹⁷ ¹⁹ ¹⁸~~56~~, ~~57~~, or ~~58~~, wherein the EPA and DHA are present in an EPA:DHA weight ratio of from 1:1 to 2:1.

²²
~~62~~. The composition of claim ²¹~~59~~, wherein the EPA and DHA are present in an EPA:DHA weight ratio of from 1:1 to 2:1.

²⁸
~~63~~. A pharmaceutical mixed-fatty-acids composition in which

Contd.
81

a) at least 80% by weight of the composition is comprised of a combination of (all-Z omega-3)-5,8,11,14,17-eicosapentaenoic acid (EPA) and (all-Z omega-3)-4,7,10,13,16,19-docosahexaenoic acid (DHA) in a weight ratio of EPA:DHA of from 1:2 to 2:1 and

b) at least 3% by weight of the composition is comprised of omega-3 fatty acids other than EPA and DHA that have 20, 21, or 22 carbon atoms.

²⁹₆₄. The composition of claim ²⁸₆₃, wherein 3 to 5% by weight of the composition is comprised of omega-3 fatty acids other than EPA and DHA that have 20, 21, or 22 carbon atoms.

³⁰₆₅. The composition of claim ²⁸₆₃, wherein at least 85% by weight of the composition is comprised of long chain omega-3 fatty acids.

³¹₆₆. The composition of claim ³⁰₆₅, wherein the EPA constitutes 40 to 60% by weight of the composition and the DHA constitutes 25 to 45% by weight of the composition.

³⁸₆₇. The composition of claim ³¹₆₆, wherein (all-Z omega-3)-6,9,12,15,18-heneicosapentaenoic acid is present in an amount of at least one percent by weight.

Contd.
B1

³²₆₈. The composition of claim ³¹₆₆, wherein C 20:4 omega-6 fatty acid constitutes at least one percent by weight of the composition.

³³₆₉. The composition of claim ³¹₆₆, wherein C 22:5 omega-3 fatty acid constitutes at least one percent by weight of the composition.

⁴⁰₇₀. The composition of claim ³⁸₆₇, wherein the (all-Z omega-3)-6,9,12,15,18-heneicosapentaenoic acid is present in an amount of from 1 to 4% by weight.

³⁴₇₁. The composition of claim ³³₆₉, wherein C 22:5 omega-3 fatty acid constitutes 1 to 3% by weight of the composition.

³¹₆₆, ⁴¹₇₂. The composition of any of claims ²⁸₆₃, ³⁰₆₅, or ³¹₆₆, wherein the EPA and DHA are present in an EPA:DHA weight ratio of from 1:1 to 2:1.

³³₆₉, ³⁹₇₃. The composition of any of claims ³⁸₆₇, ³²₆₈, or ³³₆₉, wherein the EPA and DHA are present in an EPA:DHA weight ratio of from 1:1 to 2:1.

³⁵₇₄. The composition of claim ³⁴₇₁, wherein the EPA and DHA are present in an EPA:DHA weight ratio of from 1:1 to 2:1.

³⁵₇₄, ⁴²₇₅. The composition of any of claims ²⁸₆₃, ³¹₆₆, or ³⁵₇₄, wherein the composition is in oral dosage form.

Contd.

⁴³
B1³⁵ 26. The composition of any of claims ²⁸63, ³¹66, or ³⁵74, wherein the fatty acids are present in esterified form.

⁴⁴
35 27. The composition of any of claims ²⁸63, ³¹66, or ³⁵74, wherein the fatty acids are present in ethyl ester form.

⁴⁵
35 28. The composition of any of claims ²⁸63, ³¹66, or ³⁵74, wherein the fatty acids are present in salt form.

⁴⁶
C 29. The composition of claim ²⁸74, wherein the fatty acids are present in the free acid form.

³⁷
C 30. The composition of claim ⁶⁶49, wherein the fatty acids are present in the free acid form.

³⁴
C 31. The composition of claim ⁷⁴52, wherein the fatty acids are present in the free acid form.

⁴⁷
32. A mixed-fatty-acids composition for the treatment or prophylaxis of multiple risk factors for cardiovascular diseases in which

a) at least 80% by weight of the composition is comprised of a combination of (all-Z omega-3)-

5,8,11,14,17-eicosapentaenoic acid (EPA) and (all-Z omega-3)-

4,7,10,13,16,19-docosahexaenoic acid (DHA) in a weight ratio

✓ of EPA:DHA of from 1:2 to 2:1 and

b) at least 3% by weight of the composition is comprised of omega-3 fatty acids other than EPA and DHA

C that have ²⁰18, 21, or 22 carbon atoms.

Contd.
B1

48

83. The composition of claim 82, wherein 3 to 5% by weight of the composition is comprised of omega-3 fatty acids other than EPA and DHA that have 20, 21, or 22 carbon atoms.

47

49

84. A pharmaceutical mixed-fatty-acids composition in which

a) at least 90% by weight of the composition is comprised of long chain, polyunsaturated, omega-3 fatty acids;

b) at least 80% by weight of the composition is comprised of a combination of (all-Z omega-3)-5,8,11,14,17-eicosapentaenoic acid (EPA) and (all-Z omega-3)-4,7,10,13,16,19-docosahexaenoic acid (DHA) in a weight ratio of EPA:DHA of from 1:1 to 2:1, with the EPA constituting 40 to 60% by weight of the composition and the DHA constituting 25 to 45% by weight of the composition;

c) at least 4.5% by weight of the composition is comprised of omega-3 fatty acids other than EPA and DHA that have 20, 21, or 22 carbon atoms;

d) from 1 to 4% by weight of the composition is comprised of (all-Z omega-3)-6,9,12,15,18-heneicosapentaenoic acid; and

e) the composition is in oral dosage form and includes an effective amount of a pharmaceutically acceptable antioxidant.

Contd.

81

~~50~~ ⁴⁹ 85. The composition of claim ~~84~~, wherein the fatty acids are present in ethyl ester form.

51

~~52~~ ⁵⁰ 86. The composition of claim ~~85~~, wherein at least 85% by weight of the fatty acid content of the composition is comprised of the combination of EPA and DHA.

52

~~51~~ ⁵⁰ ~~87~~ 87. The composition of either of claims ~~85~~ or ~~86~~, wherein the antioxidant is tocopherol.

24

~~53~~ 88. A mixed-fatty-acids composition for the treatment or prophylaxis of multiple risk factors for cardiovascular diseases in which

a) at least 80% by weight of the composition is comprised of omega-3 fatty acids,

b) at least 80% by weight of the total fatty acid content of the composition is comprised of a combination of (all-Z omega-3)-5,8,11,14,17-eicosapentaenoic acid (EPA) and (all-Z omega-3)-4,7,10,13,16,19-docosahexaenoic acid (DHA) in a weight ratio of EPA:DHA of from 1:2 to 2:1, and

c) omega-3 fatty acids other than EPA and DHA are present in an amount of at least 1.5% by weight of the total fatty acids.

25

~~54~~ ²⁴ ~~89~~ 89. The composition according to claim ~~88~~, wherein other long chain fatty acids present are selected from the group consisting of (all-Z omega-3)-6,9,12,15,18-heneicosapentaenoic acid, (all-Z omega-3)-7,10,13,16,19-

Contd.

~~81~~ docosapentaenoic acid, and (all-Z omega-3)-6,9,12,15-octadecatetraenoic acid.

~~25-54~~ ⁵⁶ 90. The composition according to claim ~~88~~ ^{53 24} or claim ~~89~~, wherein the total concentration of long chain omega-3 fatty acids is at least 90% by weight, the combined weight of the EPA and DHA constitutes at least 85% by weight of the total fatty acids, the EPA and DHA are present in a weight ratio of EPA:DHA of from 1:1 to 2:1, and the other long chain omega-3 C 20, C 21 and C 22 acids constitute at least 4.5% by weight of the composition.

~~51~~ ⁵⁷ 91. The composition according to claim ~~88~~ ^{53 24}, wherein the total concentration of long chain omega-3 fatty acids is at least 95% by weight, the combined weight of the EPA and DHA constitutes at least 90% by weight of the total fatty acids, and the other long chain C 20, C 21 and C 22 acids constitute at least 4.5% by weight of the composition.

~~25-54~~ ⁵⁹ 92. The composition according to claim ~~88~~ ^{53 24} or claim ~~89~~, wherein the total concentration of long chain omega-3 fatty acids is at least 85% by weight and the other long chain C 20, C 21 and C 22 acids constitute at least 4.5% by weight of the composition.

~~58~~ ⁵⁷ 93. The composition according to claim ~~91~~, wherein EPA and DHA are present in a weight ratio of EPA:DHA of from 1:1 to 2:1.

⁶⁰94. The composition according to claim ^{53 24}88, wherein the fatty acids are present in the form of pharmaceutically acceptable salts.

⁶¹95. The composition according to claim ^{53 24}88, wherein the fatty acids are present in the form of an ester.

⁶²96. The composition according to claim ⁶¹95, wherein the fatty acids are present in the form of ethyl esters.

⁶³97. The composition according to claim ⁶¹95, wherein the ester is an alkyl ester.

^{24 53}98. The composition of any of claims ^{53 24}53, ^{14 22}62, or 88, wherein the composition is in oral dosage form.

^{25 54}99. The composition of any of claims ^{54 25}53, ^{14 22}62, or 88, wherein the fatty acids are present in esterified form.

^{25 54}100. The composition of any of claims ²⁶53, ^{14 22}62, or 88, wherein the fatty acids are present in ethyl ester form.

⁵⁵101. The composition of claim ⁵⁴89, wherein the fatty acids are present in salt form.

^{25 54}102. The composition of any of claims ²⁷53, ^{14 22}62, or 89, wherein the fatty acids are present in the free acid form. --